

> d his nofil

(FILE 'HOME' ENTERED AT 09:39:49 ON 26 APR 2006)

FILE 'REGISTRY' ENTERED AT 09:40:12 ON 26 APR 2006

E 3,3,14,14-TETRAMETHYL/CN  
L1 1 SEA ABB=ON PLU=ON "3,3,14,14-TETRAMETHYLHEXADECANEDIOIC  
ACID"/CN  
D SCA  
D  
L2 STR 87272-20-6  
L3 0 SEA FAM SAM L2  
L4 1 SEA FAM FUL L2

FILE 'HCAPLUS' ENTERED AT 09:42:15 ON 26 APR 2006

L5 43 SEA ABB=ON PLU=ON L4

FILE 'MEDLINE, EMBASE, BIOSIS' ENTERED AT 09:43:02 ON 26 APR 2006

L6 65 SEA ABB=ON PLU=ON L4

FILE 'HCAPLUS' ENTERED AT 09:43:12 ON 26 APR 2006

E SYNDROME X/CT  
E E3+ALL  
E E2+ALL  
L7 2863 SEA ABB=ON PLU=ON "DISEASE, ANIMAL (L) METABOLIC SYNDROME  
X"+PFT,NT/CT  
E DYSLIPOPROTEINEM/CT  
E E4+ALL  
E E2+ALL  
L8 227 SEA ABB=ON PLU=ON "LIPOPROTEINS (L) DYSLIPOPROTEINEMIA"+PFT,N  
T/CT  
E PLASMA TRIGLYCERIDES/CT  
E E3+ALL  
E E2+ALL  
L9 20505 SEA ABB=ON PLU=ON "GLYCERIDES (L) BLOOD"+PFT,NT/CT  
E HDL CHOLESTEROL/CT  
E HDL/CT  
E E3+ALL  
E CHOLESTEROL/CT  
E E3+ALL  
E HDL/CT  
E E3+ALL  
E E2+ALL  
L10 24207 SEA ABB=ON PLU=ON "LIPOPROTEINS (L) HIGH-D."+PFT,NT/CT  
L11 208295 SEA ABB=ON PLU=ON (L7 OR L8 OR L9 OR L10) OR SYNDROM?(2A)X  
OR TRIGLYCERID? OR HDL OR CHOLESTEROL? OR DYSLIPOPROT?  
L12 19 SEA ABB=ON PLU=ON L4 AND L11

FILE 'MEDLINE' ENTERED AT 09:49:56 ON 26 APR 2006

L13 28 SEA ABB=ON PLU=ON L4  
E SYNDROME X/CT

FILE 'MEDLINE, EMBASE, BIOSIS' ENTERED AT 09:50:33 ON 26 APR 2006

L14 65 SEA ABB=ON PLU=ON L4  
L15 528315 SEA ABB=ON PLU=ON SYNDROM?(2A) X OR TRIGLYCERID? OR HDL OR  
CHOLESTEROL? OR DYSLIPOPROT? OR HIGH(3A) (D OR DENS?) (3A) (LIP?  
OR CHOLESTER?)  
L16 27 SEA ABB=ON PLU=ON L14 AND L15

=> fil hcap

FILE 'HCAPLUS' ENTERED AT 09:52:42 ON 26 APR 2006

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FILE COVERS 1907 - 26 Apr 2006 VOL 144 ISS 18

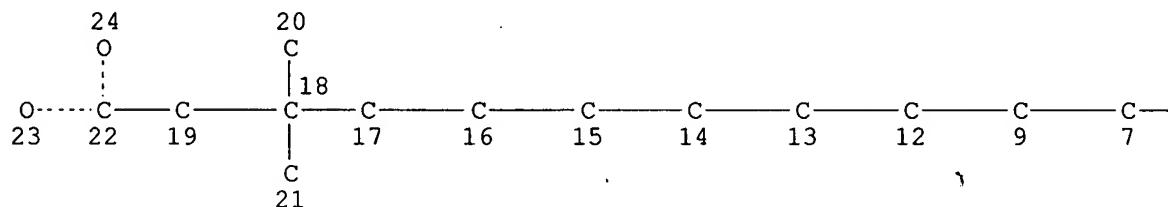
FILE LAST UPDATED: 25 Apr 2006 (20060425/ED)

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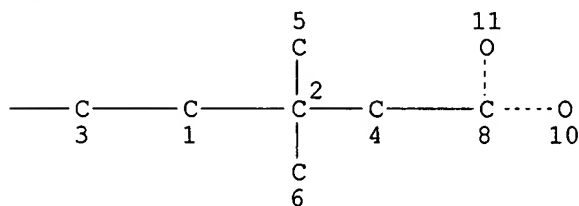
This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d que stat 112

L2 STR



Page 1-A



Page 1-B

NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 24

STEREO ATTRIBUTES: NONE

L4 1 SEA FILE=REGISTRY FAM FUL L2

L7 2863 SEA FILE=HCAPLUS ABB=ON PLU=ON "DISEASE, ANIMAL (L) METABOLIC SYNDROME X"+PFT,NT/CT

L8 227 SEA FILE=HCAPLUS ABB=ON PLU=ON "LIPOPROTEINS (L) DYSLIPOPROTEINEMIA"+PFT,NT/CT  
 L9 20505 SEA FILE=HCAPLUS ABB=ON PLU=ON "GLYCERIDES (L) BLOOD"+PFT,NT/CT  
 L10 24207 SEA FILE=HCAPLUS ABB=ON PLU=ON "LIPOPROTEINS (L) HIGH-D."+PFT,NT/CT  
 L11 208295 SEA FILE=HCAPLUS ABB=ON PLU=ON (L7 OR L8 OR L9 OR L10) OR SYNDROM?(2A)X OR TRIGLYCERID? OR HDL OR CHOLESTEROL? OR DYSLIPOPROT?  
 L12 19 SEA FILE=HCAPLUS ABB=ON PLU=ON L4 AND L11

=> fil medline embase biosis

FILE 'MEDLINE' ENTERED AT 09:52:55 ON 26 APR 2006

FILE 'EMBASE' ENTERED AT 09:52:55 ON 26 APR 2006

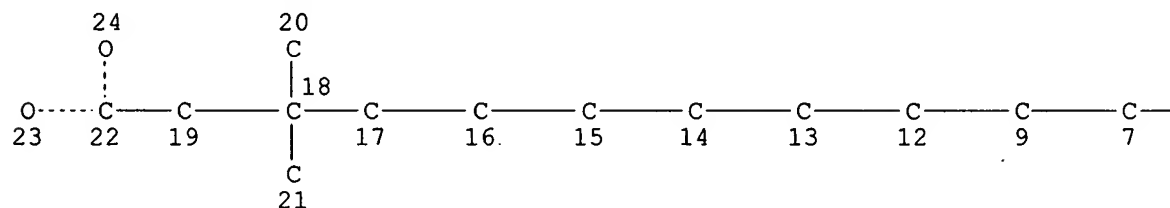
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FILE 'BIOSIS' ENTERED AT 09:52:55 ON 26 APR 2006

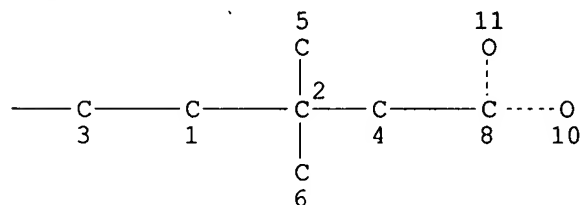
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=> d que stat l16

L2 STR



Page 1-A



Page 1-B

NODE ATTRIBUTES:

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 24

STEREO ATTRIBUTES: NONE

L4 1 SEA FILE=REGISTRY FAM FUL L2

L14 65 SEA L4

L15 528315 SEA SYNDROM?(2A) X OR TRIGLYCERID? OR HDL OR CHOLESTEROL? OR DYSLIPOPROT? OR HIGH(3A) (D OR DENS?) (3A) (LIP? OR CHOLESTER?)

L16 27 SEA L14 AND L15

=> dup rem 112 116

FILE 'HCAPLUS' ENTERED AT 09:53:03 ON 26 APR 2006  
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FILE 'BIOSIS' ENTERED AT 09:53:03 ON 26 APR 2006

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PROCESSING COMPLETED FOR L12

PROCESSING COMPLETED FOR L16

L17 26 DUP REM L12 L16 (20 DUPLICATES REMOVED)

ANSWERS '1-19' FROM FILE HCAPLUS

ANSWERS '20-21' FROM FILE MEDLINE

ANSWERS '22-26' FROM FILE EMBASE

=> d 117 ibib abs hitind hitstr 1-26

L17 ANSWER 1 OF 26 HCAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2005:1162213 HCAPLUS

DOCUMENT NUMBER: 144:408

TITLE: Transcriptional suppression of human microsomal  
**triglyceride** transfer protein by hypolipidemic  
insulin sensitizers

AUTHOR(S): Sheena, Vered; Hertz, Rachel; Berman, Ina; Noursbeck,  
Janna; Bar-Tana, Jacob

CORPORATE SOURCE: Department of Human Nutrition and Metabolism, Hebrew  
University Medical School, Jerusalem, 91120, Israel

SOURCE: Biochemical Pharmacology (2005), 70(11), 1548-1559  
CODEN: BCPA6; ISSN: 0006-2952

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Microsomal **triglyceride** transfer protein (MTP) catalyzes the  
assembly and secretion of liver **triglyceride**-rich lipoproteins.  
The human MTP (hMTP) promoter activity is reported here to be suppressed  
by HNF-4 $\alpha$  ligand antagonists (e.g., Medica analogs) or by  
PPAR $\gamma$  ligand agonists (e.g., thiazolidinediones), thus accounting  
for their hypolipidemic activity in humans. Suppression of liver hMTP by  
Medica analogs or by thiazolidinediones was mediated by the TAAA sequence  
that serves as non-canonical TATA box of the hMTP core promoter. MTP  
suppression was evident in the specific context of the wild type hMTP core  
promoter, but not in the context of the mutated rodent-conforming hMTP  
core promoter governed by a canonical TATA box conjoined with its proximal  
(-50/-38)DR-1 element. HMTF suppression by Medica analogs or  
thiazolidinediones mediated by hMTP TAAA was independent of HNF-4 $\alpha$   
or PPAR $\gamma$ . HMTF suppression by Medica analogs, but not by  
thiazolidinediones, was further complemented by inhibition of HNF-4 $\alpha$   
transcriptional activity transduced by the distal (-83/-70)DR-1 element of  
hMTP promoter. HMTF promoter activity was unaffected by PPAR $\alpha$   
activation. Furthermore, in contrast to hMTP, the promoter activity of  
the rodent-conforming hMTP was robustly activated by Wy-14,643-activated  
PPAR $\alpha$  or by thiazolidinedione-activated PPAR $\gamma$ .  
Transcriptional activation by PPAR $\alpha$  or PPAR $\gamma$  of the  
rodent-conforming, but not the wild type hMTP gene promoter, resulted from

=> d his nofil

(FILE 'HOME' ENTERED AT 11:25:02 ON 26 APR 2006)

FILE 'HCAPLUS' ENTERED AT 11:25:14 ON 26 APR 2006

E BAR-TANA/AU

E BAR TANA/AU

L1 105 SEA ABB=ON PLU=ON ("BAR TANA J"/AU OR "BAR TANA JACOB"/AU)  
E BARTANA/AU

FILE 'REGISTRY' ENTERED AT 11:26:23 ON 26 APR 2006

E 3,3,14,14/CN

E 3,3,14,14-TETRAMETHYL/CN

L2 1 SEA ABB=ON PLU=ON "3,3,14,14-TETRAMETHYLHEXADECANEDIOIC  
ACID"/CN

FILE 'HCAPLUS' ENTERED AT 11:26:57 ON 26 APR 2006

L3 35 SEA ABB=ON PLU=ON L1 AND L2

L4 20 SEA ABB=ON PLU=ON L1 AND (SYNDROM?(2A)X OR DYSLIPOPROT? OR  
HDL OR CHOLESTEROL? OR TRIGLYCERID?)

L5 15 SEA ABB=ON PLU=ON L3 AND L4

L6 40 SEA ABB=ON PLU=ON L3 OR L4

=> d 16 ibib abs hitind 1-40

L6 ANSWER 1 OF 40 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1162213 HCAPLUS

DOCUMENT NUMBER: 144:408

TITLE: Transcriptional suppression of human microsomal  
**triglyceride** transfer protein by hypolipidemic  
insulin sensitizers

AUTHOR(S): Sheena, Vered; Hertz, Rachel; Berman, Ina; Nousbeck,  
Janna; **Bar-Tana, Jacob**

CORPORATE SOURCE: Department of Human Nutrition and Metabolism, Hebrew  
University Medical School, Jerusalem, 91120, Israel

SOURCE: Biochemical Pharmacology (2005), 70(11), 1548-1559  
CODEN: BCPCA6; ISSN: 0006-2952

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Microsomal **triglyceride** transfer protein (MTP) catalyzes the assembly and secretion of liver **triglyceride**-rich lipoproteins. The human MTP (hMTP) promoter activity is reported here to be suppressed by HNF-4 $\alpha$  ligand antagonists (e.g., Medica analogs) or by PPAR $\gamma$  ligand agonists (e.g., thiazolidinediones), thus accounting for their hypolipidemic activity in humans. Suppression of liver hMTP by Medica analogs or by thiazolidinediones was mediated by the TAAA-sequence that serves as non-canonical TATA box of the hMTP core promoter. MTP suppression was evident in the specific context of the wild type hMTP core promoter, but not in the context of the mutated rodent-conforming hMTP core promoter governed by a canonical TATA box conjoined with its proximal (-50/-38)DR-1 element. hMTP suppression by Medica analogs or thiazolidinediones mediated by hMTP TAAA was independent of HNF-4 $\alpha$  or PPAR $\gamma$ . hMTP suppression by Medica analogs, but not by thiazolidinediones, was further complemented by inhibition of HNF-4 $\alpha$  transcriptional activity transduced by the distal (-83/-70)DR-1 element of hMTP promoter. hMTP promoter activity was unaffected by PPAR $\alpha$  activation. Furthermore, in contrast to hMTP, the promoter activity of the rodent-conforming hMTP was robustly activated by Wy-14,643-activated